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<110> Chernajovsky, Yuti
Dreja, Hanna Stina
Adams, Gillian
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<130> 0623.1000000

<140> US 09/756,283

<141> 2001-01-09

<160> 100

<170> PatentIn version 3.0

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Trp Leu Leu Val Leu Thr Pro Gly Pro Pro Ala Ala Gly Leu Ser Thr
                                                                                             96
                                                                                            144
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atc cgc ggc cag atc ctg tcc aag ctg cgg ctc gcc agc ccc ccg agc Ile Arg Gly Gln Ile Leu Ser Lys Leu Arg Leu Ala Ser Pro Pro Ser
                                                                                            192
cag ggg gag gtg ccg ccc ggc ccg ctg ccc gag gcc gtg ctc gcc ctg Gln Gly Glu Val Pro Pro Gly Pro Leu Pro Glu Ala Val Leu Ala Leu
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Tyr Asn Ser Thr Arg Asp Arg Val Ala Gly Glu Ser Ala Glu Pro Glu
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Pro Glu Pro Glu Ala Asp Tyr Tyr Ala Lys Glu Val Thr Arg Val Leu
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atg gtg gaa acc cac aac gaa atc tat gac aag ttc aag cag agt aca
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Met Val Glu Thr His Asn Glu Ile Tyr Asp Lys Phe Lys Gln Ser Thr
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cac agc ata tat atg ttc ttc aac aca tca gag ctc cga gaa gcg gta
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His Ser Ile Tyr Met Phe Phe Asn Thr Ser Glu Leu Arg Glu Ala Val
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Gly	Gly 290	Ser	Ala	Ala	Ala	11e 295	Asn	tat Tyr	ьys	GIII	300	GIII	цец	GIII	Giu	912
agg Arg 305	Thr	aac Asn	att Ile	cgg Arg	aaa Lys 310	tgt Cys	cag Gln	gag Glu	ctc Leu	ctg Leu 315	gag Glu	cag Gln	ctg Leu	aat Asn	gga Gly 320	960
aag Lys	atc Ile	aac Asn	ctc Leu	acc Thr 325	Tyr	agg Arg	gcg Ala	gac Asp	ttc Phe 330	ьys	atc Ile	cct Pro	atg Met	gag Glu 335	MEC	1008
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Ile Arg Gly Gln Ile Leu Ser Lys Leu Arg Leu Ala Ser Pro Pro Ser 50 55 60

Gln Gly Glu Val Pro Pro Gly Pro Leu Pro Glu Ala Val Leu Ala Leu 65 70 75 80

Tyr Asn Ser Thr Arg Asp Arg Val Ala Gly Glu Ser Ala Glu Pro Glu 85 90 95

Pro Glu Pro Glu Ala Asp Tyr Tyr Ala Lys Glu Val Thr Arg Val Leu 100 105 110

Met Val Glu Thr His Asn Glu Ile Tyr Asp Lys Phe Lys Gln Ser Thr 115 120 125

His Ser Ile Tyr Met Phe Phe Asn Thr Ser Glu Leu Arg Glu Ala Val 130 135 140

Pro Glu Pro Val Leu Leu Ser Arg Ala Glu Leu Arg Leu Leu Arg Arg 145 150 150

Leu Lys Leu Lys Val Glu Gln His Val Glu Leu Tyr Gln Lys Tyr Ser 165 170 175

Asn Asn Ser Trp Arg Tyr Leu Ser Asn Arg Leu Leu Ala Pro Ser Asp 180 185

Ser Pro Glu Trp Leu Ser Phe Asp Val Thr Gly Val Val Arg Gln Trp Leu Ser Arg Gly Gly Glu Ile Glu Gly Phe Arg Leu Ser Ala His Cys Ser Cys Asp Ser Arg Asp Asn Thr Leu Gln Val Asp Ile Asn Gly Phe Thr Thr Gly Arg Gly Asp Leu Ala Thr Ile His Gly Met Asn Arg Pro Phe Leu Leu Met Ala Thr Pro Leu Glu Arg Ala Gln His Leu Gln Ser Glu Phe Gly Gly Gly Ser Pro Leu Gly Leu Trp Ala Gly Gly Gly Ser Ala Ala Ala Ile Asn Tyr Lys Gln Leu Gln Leu Gln Glu Arg Thr Asn Ile Arg Lys Cys Gln Glu Leu Leu Glu Gln Leu Asn Gly Lys Ile Asn Leu Thr Tyr Arg Ala Asp Phe Lys Ile Pro Met Glu Met Thr Glu Lys Met Gln Lys Ser Tyr Thr Ala Phe Ala Ile Gln Glu Met Leu Gln Asn Val Phe Leu Val Phe Arg Asn Asn Phe Ser Ser Thr Gly 355 365 Trp Asn Glu Thr Ile Val Val Arg Leu Leu Asp Glu Leu His Gln Gln Thr Val Phe Leu Lys Thr Val Leu Glu Glu Lys Gln Glu Glu Arg Leu Thr Trp Glu Met Ser Ser Thr Ala Leu His Leu Lys Ser Tyr Tyr Trp Arg Val Gln Arg Tyr Leu Lys Leu Met Lys Tyr Asn Ser Tyr Ala Trp Met Val Val Arg Ala Glu Ile Phe Arg Asn Phe Leu Ile Ile Arg Arg

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m Trp}$ Glu Met Ser Ser ${
m Thr}$ Ala Leu His Leu Lys Ser ${
m Tyr}$ ${
m Trp}$ Arg 432 135 gtg caa agg tac ctt aaa ctc atg aag tac aac agc tac gcc tgg atg 480 Val Gln Arg Tyr Leu Lys Leu Met Lys Tyr Asn Ser Tyr Ala Trp Met gtg gtc cga gca gag atc ttc agg aac ttt ctc atc att cga aga ctt 528 Val Val Arg Ala Glu Ile Phe Arg Asn Phe Leu Ile Ile Arg Arg Leu 170 acc aga aac ttc caa aac gaa ttc ggg gga ggc gga tcc ccg ctc ggg Thr Arg Asn Phe Gln Asn Glu Phe Gly Gly Gly Ser Pro Leu Gly

576

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gag gcc gac Glu Ala Asp 275	tac tac g Tyr Tyr A	cc aag gag la Lys Glu 280	gtc acc Val Thr	cgc gtg cta Arg Val Leu 285	atg gtg Met Val	gaa 864 Glu
acc cac aac Thr His Asn 290	gaa atc t Glu Ile T	at gac aag yr Asp Lys 295	ttc aag Phe Lys	cag agt aca Gln Ser Thr 300	cac agc His Ser	ata 912 Ile
tat atg ttc Tyr Met Phe 305	Phe Asn T	ca tca gag nr Ser Glu 10	ctc cga Leu Arg	gaa gcg gta Glu Ala Val 315	cct gaa Pro Glu	CCC 960 Pro 320
gtg ttg ctc Val Leu Leu	tcc cgg g Ser Arg A 325	ca gag ctg la Glu Leu	cgt ctg Arg Leu 330	ctg agg agg Leu Arg Arg	ctc aag Leu Lys 335	tta 1008 Leu
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gga ggg gaa Gly Gly Glu 385	Ile Glu G	gc ttt cgc ly Phe Arc 90	ctt agc Leu Ser	gcc cac tgc Ala His Cys 395	tcc tgt Ser Cys	gac 1200 Asp 400
agc agg gat Ser Arg Asp	aac aca c Asn Thr L 405	tg caa gto eu Gln Val	g gac atc Asp Ile 410	aac ggg tto Asn Gly Phe	act acc Thr Thr 415	ggc 1248 Gly
cgc cga ggt Arg Arg Gly	gac ctg g Asp Leu A 420	cc acc att la Thr Ile	cat ggc His Gly 425	atg aac cgg Met Asn Arg	cct ttc Pro Phe 430	ctg 1296 Leu
ctt ctc atg Leu Leu Met 435	gcc acc c Ala Thr F	cg ctg gag ro Leu Glu 440	ı Arg Ala	cag cat cto Gln His Let 445	GIn Ser	tga 1344
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Thr Asn Ile Arg Lys Cys Gln Glu Leu Leu Glu Gln Leu Asn Gly Lys 35 40 45

Ile Asn Leu Thr Tyr Arg Ala Asp Phe Lys Ile Pro Met Glu Met Thr 50 55 60

Glu Lys Met Gln Lys Ser Tyr Thr Ala Phe Ala Ile Gln Glu Met Leu 65 70 75 80

Gln Asn Val Phe Leu Val Phe Arg Asn Asn Phe Ser Ser Thr Gly Trp 85 90 95

Asn Glu Thr Ile Val Val Arg Leu Leu Asp Glu Leu His Gln Gln Thr

Val Phe Leu Lys Thr Val Leu Glu Glu Lys Gln Glu Glu Arg Leu Thr 115 120 125

Trp Glu Met Ser Ser Thr Ala Leu His Leu Lys Ser Tyr Tyr Trp Arg 130 135 140

Val Gln Arg Tyr Leu Lys Leu Met Lys Tyr Asn Ser Tyr Ala Trp Met 145 150 155 160

Val Val Arg Ala Glu Ile Phe Arg Asn Phe Leu Ile Ile Arg Arg Leu 165 170 175

Thr Arg Asn Phe Gln Asn Glu Phe Gly Gly Gly Ser Pro Leu Gly 180 185 190

Leu Trp Ala Gly Gly Ser Ala Ala Leu Ser Thr Cys Lys Thr
195 200 205

Ile Asp Met Glu Leu Val Lys Arg Lys Arg Ile Glu Ala Ile Arg Gly 210 215 220

Gln Ile Leu Ser Lys Leu Arg Leu Ala Ser Pro Pro Ser Gln Gly Glu

230 225 235 240 Val Pro Pro Gly Pro Leu Pro Glu Ala Val Leu Ala Leu Tyr Asn Ser Thr Arg Asp Arg Val Ala Gly Glu Ser Ala Glu Pro Glu Pro Glu Pro Glu Ala Asp Tyr Tyr Ala Lys Glu Val Thr Arg Val Leu Met Val Glu Thr His Asn Glu Ile Tyr Asp Lys Phe Lys Gln Ser Thr His Ser Ile Tyr Met Phe Phe Asn Thr Ser Glu Leu Arg Glu Ala Val Pro Glu Pro Val Leu Leu Ser Arg Ala Glu Leu Arg Leu Leu Arg Arg Leu Lys Leu Lys Val Glu Gln His Val Glu Leu Tyr Gln Lys Tyr Ser Asn Asn Ser 340 345 350Trp Arg Tyr Leu Ser Asn Arg Leu Leu Ala Pro Ser Asp Ser Pro Glu Trp Leu Ser Phe Asp Val Thr Gly Val Val Arg Gln Trp Leu Ser Arg Gly Gly Glu Ile Glu Gly Phe Arg Leu Ser Ala His Cys Ser Cys Asp Ser Arg Asp Asn Thr Leu Gln Val Asp Ile Asn Gly Phe Thr Thr Gly Arg Arg Gly Asp Leu Ala Thr Ile His Gly Met Asn Arg Pro Phe Leu

Leu Leu Met Ala Thr Pro Leu Glu Arg Ala Gln His Leu Gln Ser 435 440 445

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<211> 390

<212> PRT

<213> Homo sapiens

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Arg Ser Cys Lys Cys Ser 385 390

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<213> Homo sapiens

<400> 24

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Asp Ala Ala Tyr Cys Phe Arg Asn Val Gln Asp Asn Cys Cys Leu Arg 305 310 315 320

Pro Leu Tyr Ile Asp Phe Lys Arg Asp Leu Gly Trp Lys Trp Ile His 325 330 335

Glu Pro Lys Gly Tyr Asn Ala Asn Phe Cys Ala Gly Ala Cys Pro Tyr 340 345 350

Leu Trp Ser Ser Asp Thr Gln His Ser Arg Val Leu Ser Leu Tyr Asn 355 360 365

Thr Glu Asn Pro Glu Ala Ser Ala Ser Pro Cys Cys Val Ser Gln Asp 370 380

Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Ile Gly Lys Ile Pro Lys Ile 385 395 400

Glu Gln Leu Ser Asn Met Ile Val Lys Ser Cys Lys Cys Ser 405 410

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<212> PRT

<213> Homo sapiens

<400> 25

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Gly His Ile Lys Lys Lys Arg Val Glu Ala Ile Arg Gly Gln Ile Leu 35 40 45

Ser Lys Leu Arg Leu Thr Ser Pro Pro Glu Pro Thr Val Met Thr His 50 60

Val Pro Tyr Gln Val Leu Ala Leu Tyr Asn Ser Thr Arg Glu Leu Leu 65 70 75 80

Glu Glu His Gly Glu Arg Lys Glu Glu Gly Cys Thr Gln Glu Asn Thr 85 90 95

Glu Ser Glu Tyr Tyr Ala Lys Glu Ile His Lys Phe Asp Met Ile Gln
100 105 110

Gly Leu Ala Glu His Asn Glu Leu Ala Val Cys Pro Lys Gly Ile Thr $115 \hspace{1.5cm} 120 \hspace{1.5cm} 125$

Ser Lys Val Phe Arg Phe Asn Val Ser Ser Val Glu Lys Asn Arg Thr 130 135 140

Asn Leu Phe Arg Ala Glu Phe Arg Val Leu Arg Val Pro Asn Pro Ser 145 150 155

Ser Lys Arg Asn Glu Gln Arg Ile Glu Leu Phe Gln Ile Leu Arg Pro 165 170 175 Asp Glu His Ile Ala Lys Gln Arg Tyr Ile Gly Gly Lys Asn Leu Pro 180 185 190 Thr Arg Gly Thr Ala Glu Trp Leu Ser Phe Asp Val Thr Asp Thr Val 200 Arg Glu Trp Leu Leu Arg Arg Glu Ser Asn Leu Gly Leu Glu Ile Ser 215 Ile His Cys Pro Cys His Thr Phe Gln Pro Asn Gly Asp Ile Leu Glu Asn Ile His Glu Val Met Glu Ile Lys Phe Lys Gly Val Asp Asn Glu Asp Asp His Gly Arg Gly Asp Leu Gly Arg Leu Lys Lys Gln Lys Asp 265 Asn Asn Asn Pro His Leu Ile Leu Met Met Ile Pro Pro His Arg Leu Asp Asn Pro Gly Gln Gly Gln Arg Lys Lys Arg Ala Leu Asp Ile 290 295 300 Asn Tyr Cys Phe Arg Asn Leu Glu Glu Asn Cys Cys Val Arg Pro Leu 305 310 315 Tyr Ile Asp Phe Arg Gln Asp Leu Gly Trp Lys Trp Val His Glu Pro 325 330 335 Lys Gly Tyr Tyr Ala Asn Phe Cys Ser Gly Pro Cys Pro Tyr Leu Arg 340 345 350Ser Ala Asp Thr Thr His Ser Thr Val Leu Gly Leu Tyr Asn Thr Leu Asn Pro Glu Ala Ser Ala Ser Pro Cys Cys Val Pro Gln Asp Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Val Gly Arg Thr Pro Lys Val Glu Gln 385 395 400 Leu Ser Asn Met Val Val Lys Ser Cys Lys Cys Ser <210> 26

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Ala Ala Ala Asp Ser Ala Gly Thr Glu Gln Arg Leu Glu Leu Tyr Gln Gly Tyr Gly Asn Ala Ser Trp Arg Tyr Leu His Gly Arg Ser Val Arg Ala Thr Ala Asp Asp Glu Trp Leu Ser Phe Asp Val Thr Asp Ala Val His Gln Trp Leu Ser Gly Ser Glu Leu Leu Gly Val Phe Lys Leu Ser Val His Cys Pro Cys Glu Met Gly Pro Gly His Ala Asp Glu Met Arg Ile Ser Ile Glu Gly Phe Glu Gln Gln Arg Gly Asp Met Gln Ser Ile Ala Lys Lys His Arg Arg Val Pro Tyr Val Leu Ala Met Ala Leu Pro Ala Glu Arg Ala Asn Glu Leu His Ser Ala Arg Arg Arg Asp Leu 180 185 190 Asp Thr Asp Tyr Cys Phe Gly Pro Gly Thr Asp Glu Lys Asn Cys Cys Val Arg Pro Leu Tyr Ile Asp Phe Arg Lys Asp Leu Gln Trp Lys Trp Ile His Glu Pro Lys Gly Tyr Met Ala Asn Phe Cys Met Gly Pro Cys 225 230 235 240 Pro Tyr Ile Trp Ser Ala Asp Thr Gln Tyr Ile Lys Val Leu Ala Leu 245 250 255 Tyr Asn Gln Asn Asn Pro Gly Ala Ser Ala Ala Pro Cys Cys Val Pro 260 265 270 Gln Ile Leu Asp Pro Leu Pro Ile Ile Tyr Tyr Val Gly Arg Asn Val 280 Arg Val Glu Gln Leu Ser Asn Met Val Val Arg Ala Cys Lys Cys Ser 290 295 <210> 27 <211> 383

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Pro Ser Glu Ala Ile Phe Leu Tyr Asn Ser Thr Leu Glu Val Ile Arg Glu Lys Ala Thr Arg Glu Glu Glu Glu His Val Gly His Asp Gln Asn Ile Gln Asp Tyr Tyr Ala Lys Gln Val Tyr Arg Phe Glu Ser Ile Thr Glu Leu Glu Asp His Glu Phe Lys Phe Lys Phe Asn Ala Ser Asn Val Arg Glu Asn Val Gly Met Asn Ser Leu Leu His His Ala Glu Leu Arg 135 Met Tyr Lys Lys Gln Thr Asp Lys Asn Met Asp Gln Arg Met Glu Leu Phe Trp Lys Tyr Gln Glu Asn Gly Thr Thr His Ser Arg Tyr Leu Glu Ser Lys Tyr Ile Thr Pro Val Thr Asp Asp Glu Trp Met Ser Phe Asp 185 Val Thr Lys Thr Val Asn Glu Trp Leu Lys Arg Ala Glu Glu Asn Glu Gln Phe Gly Leu Gln Pro Ala Cys Lys Cys Pro Thr Pro Gln Ala Lys Asp Ile Asp Ile Glu Gly Phe Pro Ala Leu Arg Gly Asp Leu Ala Ser Leu Ser Ser Lys Glu Asn Thr Lys Pro Tyr Leu Met Ile Thr Ser His Pro Ala Glu Arg Ile Asp Thr Val Thr Ser Ser Arg Lys Lys Arg Gly 265 Val Gly Gln Glu Tyr Cys Phe Gly Asn Asn Gly Pro Asn Cys Cys Val Lys Pro Leu Tyr Ile Asn Phe Arg Lys Asp Leu Gly Trp Lys Trp Ile 295 His Glu Pro Lys Gly Tyr Glu Ala Asn Tyr Cys Leu Gly Asn Cys Pro Tyr Ile Trp Ser Met Asp Thr Gln Tyr Ser Lys Val Leu Ser Leu Tyr Asn Gln Asn Asn Pro Gly Ala Ser Ile Ser Pro Cys Cys Val Pro Asp 345 Val Leu Glu Pro Leu Pro Ile Ile Tyr Tyr Val Gly Arg Ile Ala Lys Val Glu Gln Leu Ser Asn Met Val Val Arg Ser Cys Asn Cys Ser 375

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